

FORM PTO-1449
THIRD SUPPLEMENTAL
INFORMATION DISCLOSURE STATEMENT

ATTY. DOCKET NO.
1503.1030002/JMC/THN/JSL

APPLICATION NO.
10/058,215

APPLICANT
Lu et al.

FILING DATE
January 29, 2002

GROUP
1614

U.S. PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB- CLASS	FILING DATE
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FOREIGN PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB- CLASS	TRANSLATION
	AL						Yes No
	AM						Yes No
	AN						Yes No
KH	AO3	EP 0 623 607 A1	11/09/1994	Europe			Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

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DATE CONSIDERED

Aug. 27, 2003

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KN	AI1	6,066,648	05/23/2000	Duggan et al.	514	300	12/15/1998
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FOREIGN PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB- CLASS	TRANSLATION
	AL						Yes No
	AM						Yes No
KN	AN3	WO 00/02874	01/20/2000	WIPO			Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

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KN	AA1	5,741,796	04/21/1998	Hartman et al.	514	300	09/26/1996
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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB- CLASS	TRANSLATION
KN	AL1	WO 97/06791	02/27/1997	WIPO			Yes No
I	AM1	WO 97/23451	07/03/1997	WIPO			X Yes No
	AN1	WO 97/36859	10/09/1997	WIPO			Yes No
	AO1	WO 97/45137	12/04/1997	WIPO			Yes No
↓	AP1	WO 98/00395	01/08/1998	WIPO			Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

KN	AR	1	Albelda, S.M., "Biology of Disease. Role of Integrins and Other Cell Adhesion Molecules in Tumor Progression and Metastasis," Lab. Invest. 68:4-17, The United States and Canadian Academy of Pathology, Inc. (1993).
I	AS	1	Albelda, S.M. et al., "Integrin Distribution in Malignant Melanoma: Association of the β_3 Subunit with Tumor Progression and Metastasis," Cancer Res. 50:6757-6764, American Association for Cancer Research (1990).
↓	AT	1	Batcho, A. et al., "Indoles From 2-Methylnitrobenzenes by Condensation With Formamide Acetals Followed by Reduction: 4-Benzylloxyindole (1 H-Indole, 4-(phenylmethoxy)-)," in Organic Synthesis, Saucy, G. et al., Eds., John Wiley & Sons, New York, NY, pp. 214-225 (1985).

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KH	AL2	WO 98/14192	04/09/98	WIPO		Yes No
KA	AM2	WO 00/33838	06/15/00	WIPO		Yes No
	AN2					Yes No
	AO2					Yes No
	AP2					Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

KH	AR	2	Boudreau, N. and Rabinovitch, M., "Developmentally Regulated Changes in Extracellular Matrix in Endothelial and Smooth Muscle Cells in the Ductus Arteriosus May be Related to Intimal Proliferation," Lab. Invest. 64:187-199, The United States and Canadian Academy of Pathology, Inc. (1991).
I	AS	2	Brocke, S. et al., "Antibodies to CD44 and integrin α_4 , but not L-selectin, prevent central nervous system inflammation and experimental encephalomyelitis by blocking secondary leukocyte recruitment," Proc. Natl. Acad. Sci. USA 96:6896-6901, The National Academy of Sciences (1999).
↓	AT	2	Brooks, P.C. et al., "Integrin $\alpha_v\beta_3$ Antagonists Promote Tumor Regression by Inducing Apoptosis of Angiogenic Blood Vessels," Cell 79:1157-1164, Cell Press (1994).

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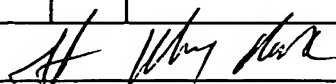
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	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

KN ↓	AR	<u>3</u>	Brooks, P.C., "Integrin $\alpha\beta$ 3: A Therapeutic Target," <i>DN&P</i> 10:456-461, Prous Science (1997).
	AS	<u>3</u>	Brooks, P.C. et al., "Antiintegrin $\alpha\beta$ 3 blocks human breast cancer growth and angiogenesis in human skin," <i>J. Clin. Invest.</i> 96:1815-1822, The American Society for Clinical Investigation, Inc. (1995).
	AT	<u>3</u>	Brooks, P.C. et al., "Requirement of Vascular Integrin $\alpha\beta$ 3 for Angiogenesis," <i>Science</i> 264:569-571, American Association for the Advancement of Science (1994).

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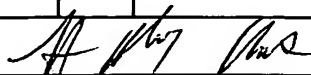
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	AL					Yes No
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	AN					Yes No
	AO					Yes No
	AP					Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

K H ↓	AR	4	Brooks, P.C., "Cell adhesion molecules in angiogenesis," <i>Cancer Met. Rev.</i> 15:187-194, Kluwer Academic Publishers (1996).
	AS	4	Cheresh, D.A., "Structure, function and biological properties of integrin of $\alpha_v\beta_3$ on human melanoma cells," <i>Cancer Met. Rev.</i> 10:3-10, Kluwer Academic Publishers (1991).
	AT	4	Choi, E.T. et al., "Inhibition of neointimal hyperplasia by blocking $\alpha_v\beta_3$ integrin with a small peptide antagonist GpenGRGDSPCA," <i>J. Vasc. Surg.</i> 19:125-134, The Society for Vascular Surgery and International Society for Cardiovascular Surgery (1994).

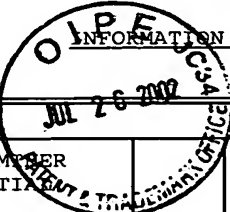
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	AN					Yes No	
	AO					Yes No	
	AP					Yes No	

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
KH	AR	<u>5</u>	Corey, E.J. and Fuchs, P.L., "A Synthetic Method for Formyl \rightarrow Ethynyl Conversion (RCHO \rightarrow RC=CH or RC=CR')," <i>Tetrahedron Lett.</i> 36:3769-3772, Pergamon Press (1972).
I	AS	<u>5</u>	Enenstein, J. and Kramer, R.H., "Confocal Microscopic Analysis of Integrin Expression on the Microvasculature and its Sprouts in the Neonatal Foreskin," <i>J. Invest. Dermatol.</i> 103:381-386, The Society for Investigative Dermatology, Inc. (1994).
✓	AT	<u>5</u>	Fisher, J.E. et al., "Inhibition of Osteoclastic Bone Resorption In Vivo by Echistatin, an "Arginyl-Glycyl-Aspartyl" (RGD)-Containing Protein," <i>Endocrinology</i> 132:1411-1413, The Endocrine Society (1993).

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	AM					Yes No
	AN					Yes No
	AO					Yes No
	AP					Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

K9	AR	6	Friedlander, M. et al., "Definition of Two Angiogenic Pathways by Distinct α_v Integrins," <i>Science</i> 270:1500-1502, American Association for the Advancement of Science (1995).
L	AS	6	Gladson, C.L., "Expression of Integrin $\alpha v \beta 3$ in Small Blood Vessels of Glioblastoma Tumors," <i>J. Neuropathol. Exp. Neurol.</i> 55:1143-1149, The American Association of Neuropathologists (1996).
L	AT	6	Greene, T.W. and Wuts, P.G.M., "Special -NH Protective Groups. Protection for the Amino Group: Special -NH Protective Groups," in <i>Protective Groups in Organic Synthesis</i> , 2nd edition, John Wiley and Sons, Inc., New York, pp. 267-287 and p. 331 (1991).

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A. J. M. R. R.

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	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
KM ↓	AR	7	Greenspoon, N. et al., "Structural Analysis of Integrin Recognition and the Inhibition of Integrin-Mediated Cell Functions by Novel Nonpeptidic Surrogates of the Arg-Gly-Asp Sequence," <i>Biochemistry</i> 32:1001-1008, American Chemical Society (1993).
	AS	7	Hardan, I. et al., "Inhibition of Metastatic Cell Colonization in Murine Lungs and Tumor-Induced Morbidity by Non-Peptidic Arg-Gly-Asp Mimetics," <i>Int. J. Cancer</i> 55:1023-1028, Wiley-Liss, Inc. (1993).
	AT	7	Hendrickson, J.B. and Hussoin, Md. S., "Facile Dehydration of Activated Ketones to Alkynes," <i>Synthesis</i> 3:217-221, Thieme Medical Publishers, Inc. (1989).

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	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
<div style="font-size: 24px; font-weight: bold;">21</div>	AR	8	Hershkoviz, R. et al., "Inhibition of CD4 ⁺ T lymphocyte binding to fibronectin and immune-cell accumulation in inflammatory sites by non-peptidic mimetics of Arg-Gly-Asp," <i>Clin. Exp. Immunol.</i> 95:270-276, Blackwell Scientific Publishers (1994).
<div style="font-size: 24px; font-weight: bold;">↓</div>	AS	8	Horton, M., "Current Status Review. Vitronectin receptor: tissue specific expression or adaption to culture?," <i>Int. J. Exp. Pathol.</i> 71:741-759, Blackwell Scientific Publications (1990).
<div style="font-size: 24px; font-weight: bold;">↓</div>	AT	8	Hynes, R.O., "Integrins: Versatility, Modulation, and Signaling in Cell Adhesion," <i>Cell</i> 69:11-25, Cell Press (1992).

EXAMINER <i>[Signature]</i>	DATE CONSIDERED <i>Aug. 27, 2003</i>
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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
KN ↓	AR AS AT	2 2 2	Juliano, R., "Signal transduction by integrins and its role in the regulation of tumor growth," <i>Cancer Met. Rev.</i> 13:25-30, Kluwer Academic Publishers (1994). Kaul, D.K. et al., "Monoclonal antibodies to $\alpha v \beta 3$ (7E3 and LM609) inhibit sickle red blood cell-endothelium interactions induced by platelet-activating factor," <i>Blood</i> 95:368-374, The American Society of Hematology (January 2000). Ku, T.W. et al., "Direct Design of a Potent Non-Peptide Fibrinogen Receptor Antagonist Based on the Structure and Conformation of a Highly Constrained Cyclic RGD Peptide," <i>J. Amer. Chem. Soc.</i> 115:8861-8862, American Chemical Society (1993).

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K1	AR	10	Marquardt, D. W., "An Algorithm for Least-Squares Estimation of Nonlinear Parameters," J. Soc. Indust. Appl. Math. 11:431-441, Society for Industrial and Applied Mathematics (1963).
	AS	10	Mitsunobu, O., "The Use of Diethyl Azodicarboxylate and Triphenylphosphine in Synthesis and Transformation of Natural Products," Synthesis 1:1-28, Georg Thieme Verlag (1981).
	AT	10	Nicosia, R.F. and Madri, J.A., "The Microvascular Extracellular Matrix. Developmental Changes During Angiogenesis in the Aortic Ring-Plasma Clot Model," Amer. J. Pathol. 128:78-90, American Association of Pathologists (1987).

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KN	AR	11	Niiya, K. et al., "Increased Surface Expression of the Membrane Glycoprotein IIb/IIIa Complex Induced by Platelet Activation. Relationship to the Binding of Fibrinogen and Platelet Aggregation," <i>Blood</i> 70:475-483, Grune & Stratton, Inc. (1987).
	AS	11	Nip, J. et al., "Coordination Expression of the Vitronectin Receptor and the Urokinase-type Plasminogen Activator Receptor in Metastatic Melanoma Cells," <i>J. Clin. Invest.</i> 95:2096-2103, The American Society for Clinical Investigation, Inc. (1995).
	AT	11	Okada, Y. et al., "Integrin $\alpha_v\beta_3$ Is Expressed in Selected Microvessels after Focal Cerebral Ischemia," <i>Amer. J. Pathol.</i> 149:37-44, American Society for Investigative Pathology (1996).

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K1	AR	12	Relton, J.K. et al., "Inhibition of $\alpha 4$ Integrin Protects Against Transient Focal Cerebral Ischemia in Normotensive and Hypertensive Rats," <i>Stroke</i> 32:199-205, American Heart Association, Inc. (January 2001).
I	AS	12	Ruoslahti, E. and Giancotti, F.G., "Integrins and Tumor Cell Dissemination," <i>Cancer Cells</i> 1:119-126, Cold Spring Harbor Laboratory Press (1989).
J	AT	12	Ruoslahti, E. and Reed, J.C., "Anchorage Dependence, Integrins, and Apoptosis," <i>Cell</i> 77:477-478, Cell Press (1994).

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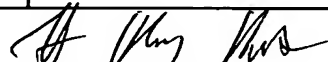
EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to Applicant.

 FORM PTO-1449 INFORMATION DISCLOSURE STATEMENT	ATTY. DOCKET NO. 1503.1030002	APPLICATION NO. 10/058,215
	APPLICANT Lu et al.	
	FILING DATE January 29, 2002	GROUP 1614

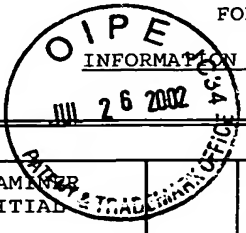
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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
	AR	<u>13</u>	Savelon, L. et al., "Substituted pyrido[3,2-b]oxazin-3(4H)-ones: synthesis and evaluation of antinociceptive activity," <i>Bioorganic Med. Chem.</i> 6:133-142, Elsevier Science Ltd. (1998).
	AS	<u>13</u>	Sato, M. et al., "Echistatin Is a Potent Inhibitor of Bone Resorption in Culture," <i>J. Cell Biol.</i> 111:1713-1723, The Rockefeller University Press (1990).
	AT	<u>13</u>	Shattil, S.J., "Function and Regulation of the B ₂ Integrins in Hemostasis and Vascular Biology," <i>Thromb. Haemost.</i> 74:149-155, F.K. Schattauer Verlagsgesellschaft mbH (1995).

EXAMINER	DATE CONSIDERED
	Aug. 27, 2007

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 <p>FORM PTO-1449 INFORMATION DISCLOSURE STATEMENT</p>		ATTY. DOCKET NO. 1503.1030002		APPLICATION NO. 10/058,215	
		APPLICANT Lu et al.			
		FILING DATE January 29, 2002		GROUP 1614	


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<div style="font-size: 2em;">K</div> <div style="font-size: 4em;">J</div>	AR	14	Sonogashira, K. et al., "A Convenient Synthesis of Acetylenes: Catalytic Substitutions of Acetylenic Hydrogen with Bromoalkenes, Iodoarenes, and Bromopyridines," <i>Tetrahedron Lett.</i> 50:4467-4470, Pergamon Press (1975).
	AS	14	Topol, E.J. et al., "Randomised trial of coronary intervention with antibody against platelet IIb/IIIa integrin for reduction of clinical restenosis: results at six months," <i>Lancet</i> 343:881-886, The Lancet Ltd. (1994).
	AT	14	White, J.M., "Integrins as virus receptors," <i>Curr. Biol.</i> 3:596-599, Current Biology (1993).

EXAMINER <i>At My Plus</i>	DATE CONSIDERED <i>Aug. 27, 2003</i>
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	APPLICANT Lu et al.	
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U.S. PATENT DOCUMENTS

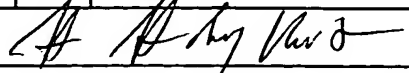
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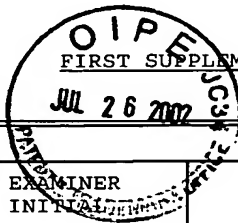
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	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

  	AR	15	Yun, Z. et al., "Involvement of Integrin $\alpha_5\beta_1$ in Cell Adhesion, Motility, and Liver Metastasis of Murine RAW117 Large Cell Lymphoma," Cancer Res. 56:3103-3111, The American Association for Cancer Research (1996).
	AS	15	Chemical Abstracts English language summary of document AM2 (WO 98/00395), CAPLUS Accession Number 1998:38464.
	AT		

EXAMINER 	DATE CONSIDERED Aug. 27, 2003
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FORM PTO-1449				ATTY. DOCKET NO. 1503.1030002/JMC/THN		APPLICATION NO. 10/058,215	
				APPLICANT Lu et al.			
				FILING DATE January 29, 2002		GROUP 1614	
U.S. PATENT DOCUMENTS							
EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB- CLASS	FILING DATE
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<i>KN</i> ↓	AB1	5,223,517	06/29/1993	Müller et al.	514	339	08/23/1991
	AC1	5,234,942	08/10/1993	Bernstein et al.	514	415	12/17/1990
	AD1	5,290,788	03/01/1994	Stevens et al.	514	314	04/21/1992
	AE1	5,744,488	04/28/1998	Cross et al.	514	339	02/23/1996
	AF1	5,834,454	11/10/1998	Kitano et al.	514	183	01/28/1997
	AG1	5,977,100	11/02/1999	Kitano et al.	514	214	05/08/1998
	AH1	6,184,238 B1	02/06/2001	Takano et al.	514	339	06/23/1999
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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB- CLASS	TRANSLATION
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	AM						Yes No
<i>K9</i> ↓	AN2	WO 98/13368 A1	04/02/1998	WIPO			Yes No
	AO2	WO 99/30713 A1	06/24/1999	WIPO			Yes No
	AP2	WO 99/32150 A1	07/01/1999	WIPO			Yes No
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)							
<i>KN</i> ↓	AR	<u>16</u>	Dennis, M.S., et al., "Binding Interactions of Kistrin With Platelet Glycoprotein IIb-IIIa: Analysis by Site-Directed Mutagenesis," <i>Proteins</i> 15:312-321, Wiley-Liss (1993)				
	AS	<u>16</u>	Fabry, M.E., et al., "Second generation knockout sickle mice: the effect of HbF," <i>Blood</i> 97:410-418, American Society of Hematology (January 2001)				
	AT	<u>16</u>	United States Pharmacopeia/ The National Formulary for 1995, published by United States Pharmacopeial Convention, Inc., Rockville, Maryland, p. 1636 (1994)				
EXAMINER <i>At King Park</i>					DATE CONSIDERED <i>Aug. 27, 2003</i>		
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FORM PTO-1449 FIRST SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT				ATTY. DOCKET NO. 1503.1030002/JMC/THN		APPLICATION NO. 10/058,215	
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				FILING DATE January 29, 2002		GROUP 1614	
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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB-CLASS	FILING DATE
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KN	AM3	WO 01/58893 A2	08/16/2001	WIPO			Yes No X
	AN						Yes No
	AO						Yes No
	AP						Yes No
OTHER (Including Author, Title, Date, Pertinent Pages, etc.)							
KH	AR	17	Dialog File 351, Accession No. 14058279, Derwent WPI English language abstract for WO 01/58893 A2 (Document AM3)				
	AS						
	AT						
EXAMINER					DATE CONSIDERED		
					Aug. 27, 2003		
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